



RECEIVED

MAR 22 2002

RECEIVED

SEQUENCE LISTING TECH CENTER 1600/2900 MAR 21 2002

TECH CENTER 1600/2900

<110> Lechler, Robert I.
Dorling, Anthony

<120> IMMUNOSUPPRESSION BY BLOCKING T CELL CO-STIMULATION SIGNAL 2 (B7/CD28 INTERACTION)

<130> 2292/OH795

<140> US 09/674,462

<141> 2001-05-08

<150> PCT/ GB99/01350

<151> 1999-04-30

<160> 27

<170> PatentIn Ver. 2.1

<210> 1

<211> 223

<212> PRT

<213> Sus scrofa

<400> 1

Met Ala Cys Ser Gly Phe Arg Ser His Gly Ala Trp Leu Glu Leu Thr
1 5 10 15

Ser Arg Thr Trp Pro Cys Thr Ala Leu Phe Ser Leu Leu Phe Ile Pro
20 25 30

Val Phe Ser Lys Gly Met His Val Ala Gln Pro Ala Val Val Leu Ala
35 40 45

Asn Ser Arg Gly Val Ala Ser Phe Val Cys Glu Tyr Gly Ser Ala Gly
50 55 60

Lys Ala Ala Glu Val Arg Val Thr Val Leu Arg Arg Ala Gly Ser Gln
65 70 75 80

Met Thr Glu Val Cys Ala Ala Thr Tyr Thr Val Glu Asp Glu Leu Thr
85 90 95

Phe Leu Asp Asp Ser Thr Cys Thr Gly Thr Ser Thr Glu Asn Lys Val
100 105 110

Asn Leu Thr Ile Gln Gly Leu Arg Ala Val Asp Thr Gly Leu Tyr Ile
115 120 125

Cys Lys Val Glu Leu Leu Tyr Pro Pro Pro Tyr Tyr Val Gly Met Gly
 130 135 140

Asn Gly Thr Gln Ile Tyr Val Ile Asp Pro Glu Pro Cys Pro Asp Ser
 145 150 155 160

Asp Phe Leu Leu Trp Ile Leu Ala Ala Val Ser Ser Gly Leu Phe Phe
 165 170 175

Tyr Ser Phe Leu Ile Thr Ala Val Ser Leu Ser Lys Met Leu Lys Lys
 180 185 190

Arg Ser Pro Leu Thr Thr Gly Val Tyr Val Lys Met Pro Pro Thr Glu
 195 200 205

Pro Glu Cys Glu Lys Gln Phe Gln Pro Tyr Phe Ile Pro Ile Asn
 210 215 220

<210> 2
 <211> 672
 <212> DNA
 <213> Sus scrofa

<400> 2
 atggcttgct ctggattccg gagccatggg gcttggtctg agcttacttc taggacctgg 60
 ccctgtacag ctctgttttc tcttctcttc atccctgtct tctccaaagg gatgcacgtg 120
 gccaacacctg cagtagtgct ggccaacagc cggggtgttg ccagctttgt gtgtgagtat 180
 gggctctgag gcaaagctgc cgaggtcccg gtgacagtgc tgcggcgggc cggcagccag 240
 atgactgaag tctgtgccgc gacataact gtggaggatg agttgacctt ccttgatgac 300
 tctacatgca ctggcacctc caccgaaaac aaagtgaacc tcaccatcca agggctgaga 360
 gccgtggaca ctgggctcta catctgcaag gtggagctcc tgtaccacc accctactat 420
 gtgggtatgg gcaacgggac ccagatttat gtcattgac cagaaccatg cccagattct 480
 gatttcctgc tctggatcct ggcagcagtt agttcagggt tgttttttta cagcttcctc 540
 atcacagctg tttctttgag caaaatgcta aagaaaagaa gtcctcttac tacaggggtc 600
 tatgtgaaaa tgcccccgac agagccagaa tgtgaaaagc aatttcagcc ttattttatt 660
 cccatcaatt ga 672

<210> 3
 <211> 400
 <212> PRT
 <213> Artificial Sequence

<220>
 <223> pCTLA4-Ig construct (Figure 4)

<400> 3
 Met Ala Cys Ser Gly Phe Arg Ser His Gly Ala Trp Leu Glu Leu Thr
 1 5 10 15

Ile Ser Lys Ala Lys Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr Leu
 290 295 300
 Pro Pro Ser Arg Asp Glu Leu Thr Lys Asn Gln Val Ser Leu Thr Cys
 305 310 315 320
 Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Glu Ser
 325 330 335
 Asn Gly Gln Pro Glu Asn Asn Tyr Lys Thr Thr Pro Pro Val Leu Asp
 340 345 350
 Ser Asp Gly Ser Phe Phe Leu Tyr Ser Lys Leu Thr Val Asp Lys Ser
 355 360 365
 Arg Trp Gln Gln Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala
 370 375 380
 Leu His Asn His Tyr Thr Gln Lys Ser Leu Ser Leu Ser Pro Gly Lys
 385 390 395 400

<210> 4
 <211> 722
 <212> DNA
 <213> Phage library

<400> 4
 ccgaggtgca gctggtggag tctgggggag gcttggtaca gcctgggggg tccctgagac 60
 tctcctgtgc agcctctgga ttcaccttta gcagctatgc catgagctgg gtccgccagg 120
 ctccaggga ggggctggag tgggtctcag ctattcgtgg tagtggtggg agcacatact 180
 acgcagactc cgtgaagggc cggttcacca tctccagaga caattccaag aacacgctgt 240
 atctgcaa at gaacagcctg agagccgagg acacggccgt gtattactgt gcaagagctg 300
 gtcgtat ttt gtttgactat tggggccaag gtaccctggg caccgtctcg agtggtggag 360
 gcggttcagg cggaggtggc tctggcggta gtgcacttca gtctgtgctg actcagccac 420
 cctcagcgtc tgggaccccc gggcagcggg tcaccatctc ttgttctgga agcagctcca 480
 acatcggaag taattatgta tactgggtacc agcagctccc aggaacggcc cccaaactcc 540
 tcatctatag gaataatcag cggccctcag gggctccctga ccgattctct ggctccaagt 600
 ctggcacctc agcctccctg gccatcagtg ggctccgggc cgaggatgag gctgattatt 660
 actgtgcagc atgggatgac agcctgggtat tcggcggagg gaccaagctc accgtcctag 720
 gt 722

<210> 5
 <211> 240
 <212> PRT
 <213> Phage library

<210> 6
<211> 729
<212> DNA
<213> Phage library

<400> 6
catggccgag gtgcagctgg tggagtctgg gggaggcttg gtacagcctg ggggggtccct 60
gagactctcc tgtgcagcct ctggattcac ctttagcagc tatgccatga gctgggtccg 120
ccaggctcca ggaagggggc tggagtgggt ctacagctatt agtggtagtg gtggtagcac 180
atactacgca gactccgtga agggccgggt caccatctcc agagacaatt ccaagaacac 240
gctgtatctg caaatgaaca gcctgagagc cgaggacacg gccgtgtatt actgtgcaag 300
agctggctcg attttggttg actattgggg ccaaggtacc ctggtcaccg tctcgagtgg 360
tggaggcggg tcaggcggag gtggctctgg cggtagtgca cttcagctctg tgctgactca 420
gccaccctca gcgtctggga cccccgggca gaggggtcacc atctcttggt ctggaagcag 480
ctccaacatc ggaagtaatt atgtatactg gtaccagcag ctcccaggaa cggcccccaa 540
actcctcatc tataggaata atcagcggcc ctccagggtc cctgaccgat tctctggctc 600
caagtctggc acctcagcct ccctggccat cagtgggctc cggtcaggag atgaggctga 660
ttattactgt gcagcatggg atgacagcct ggtattcggc ggagggacca agctgaccgt 720
cctaggtgc 729

<210> 7
<211> 738
<212> DNA
<213> Phage library

<400> 7
catggccgag gtgcagctgc aggagtcggg cccaggactg gtgaagcctc gggagaccct 60
gtccctcacc tgcactgtct ctgggtggctc cgtcagcagt ggtagttact ggagctggat 120
ccggcagccc ccagggaagg gactggagtg gattgggtat atctattaca gtgggagcac 180
caactacaac ccctccctca agagtcgagt caccatatca gtagacacgt ccaagaacca 240
gttctccctg aagctgagct ctgtgaccgc tgcggacacg gccgtgtatt actgtgcaag 300
aatgcggaag gataagtttg actattgggg ccaaggtacc ctggtcaccg tctcgagtgg 360
tggaggcggg tcaggcggag gtggctctgg cggtagtgca cttcagctctg tgctgactca 420
gccaccctca gcgtctggga cccccgggca gaggggtcacc atctcttggt ctggaagcag 480
ctccaacatc ggaagtaatt atgtatactg gtaccagcag ctcccaggaa cggcccccaa 540
actcctcatc tataggaata atcagcggcc ctccagggtc cctgaccgat tctctggctc 600
caagtctggc acctcagcct ccctggccat cagtgggctc cggtcaggag atgaggctga 660
ttattactgt gcagcatggg atgacagcct gtttgtattc ggcggaggga ccaagctgac 720
cgtccctaggg gcggccgc 738

<210> 8
<211> 739
<212> DNA
<213> Phage library

<400> 8
catggccgag gtgcagctgg tgcagtctgg ggctgagtga agaggccggg ggcctcagtg 60
aaggtttcct gcaaggcatc tggatacacc ttcaccagct actatatgca ctgggtgcga 120
caggcccctg gacaagggct tgagtggatg ggaataatca accctagtgg tggtagcaca 180

```

caagctacgc acagaagttc cagggcagag tcaccatgac cagggacacg tccacgagca 240
cagtctacat ggagctgagc agcctgagat ctgaggacac ggccgtgtat tactgtgcaa 300
gaatggctcc ctatgtgaat acgcttgttt tttggggcca aggtaccctg gtcaccgtct 360
cgagtgggtg aggcggttca ggcggagggtg gctctggcgg tagtgactt cagtctgtgc 420
tgactcagga cctgctgtg tctgtggcct tgggacagac agtcaggatc acatgccaaag 480
taggagacag cctcagaagc tattatgcaa gctggtacca gcagaagcca ggacaggccc 540
ctgtacttgt catctatggt aaaaacaacc ggccctcagg gatcccagac cgattctctg 600
gctccagctc aggaacacac gcttccttga ccatcactgg ggctcaggcg gaagatgagg 660
ctgactatta ctgtaactcc cgggacagca gtggttttac tgtattcggc ggagggacca 720
agctgaccgt cctaggtgc                                     739

```

<210> 9
 <211> 729
 <212> DNA
 <213> Phage library

```

<400> 9
catgggccc ggtgcagctg ttgcagtctg cagcagaggt gaaaaagccc ggggagtctc 60
tgaagatctc ctgtaagggt tctggataca gctttaccag ctactggatc ggctgggtgc 120
gccagatgcc cgggaaaggc ctggagtgga tggggatcat ctatcctggg gactctgata 180
ccagatacag cccgtccttc caaggccagg tcaccatctc agccgacaag tccatcagca 240
ccgcctacct gcagtggagc agcctgaagg cctcggacac ggccgtgtat tactgtgcaa 300
gattttcgct tgggtggttt gactattggg gccaaaggta cctggtcacc gtctcgagtgc 360
gtggaggcgg ttcaggcgga ggtggctctg gcggtagtgc acttgacatc cagttgacct 420
agtctccatg ttctgtctg catctgtagg agacagagtc accatcactt gccggggccag 480
tcagggcatt agcagttatt tagcctggtg tcagcaaaaa ccagggaaag cccctaagct 540
cctggtctat gctgcatcca ctttgcaaag tgggggtccc tcaagggttc gcggcagtgg 600
atctgggaca gaattcactc tcacaatcag cagcctgcag cctgaagatt ttgcaactta 660
ttactgtcaa cagcttaata gttaccgctt gacgttcggc caagggacca agctggaaat 720
caaacgtgc                                     729

```

<210> 10
 <211> 240
 <212> PRT
 <213> Phage library

```

<400> 10
Glu Val Gln Leu Val Glu Ser Gly Gly Gly Leu Val Gln Pro Gly Gly
  1              5              10              15

Ser Leu Arg Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Ser Tyr
  20              25              30

Ala Met Ser Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp Val
  35              40              45

Ser Ala Ile Ser Gly Ser Gly Gly Ser Thr Tyr Tyr Ala Asp Ser Val
  50              55              60

Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ser Lys Asn Thr Leu Tyr

```


<400> 13

Gln	Val	Gln	Leu	Leu	Gln	Ser	Ala	Ala	Glu	Val	Lys	Lys	Pro	Gly	Glu
1				5					10					15	
Ser	Leu	Lys	Ile	Ser	Cys	Lys	Gly	Ser	Gly	Tyr	Ser	Phe	Thr	Ser	Tyr
			20					25					30		
Trp	Ile	Gly	Trp	Val	Arg	Gln	Met	Pro	Gly	Lys	Gly	Leu	Glu	Trp	Met
		35					40					45			
Gly	Ile	Ile	Tyr	Pro	Gly	Asp	Ser	Asp	Thr	Arg	Tyr	Ser	Pro	Ser	Phe
	50					55					60				
Gln	Gly	Gln	Val	Thr	Ile	Ser	Ala	Asp	Lys	Ser	Ile	Ser	Thr	Ala	Tyr
65					70					75					80
Leu	Gln	Trp	Ser	Ser	Leu	Lys	Ala	Ser	Asp	Thr	Ala	Val	Tyr	Tyr	Cys
				85					90					95	
Ala	Arg	Phe	Ser	Leu	Gly	Gly	Phe	Asp	Tyr	Trp	Gly	Gln	Gly	Thr	Leu
			100					105					110		
Val	Thr	Val	Ser	Ser	Gly	Gly	Gly	Gly	Ser	Gly	Gly	Gly	Gly	Ser	Gly
		115					120					125			
Gly	Ser	Ala	Leu	Asp	Ile	Gln	Leu	Thr	Gln	Ser	Pro	Ser	Phe	Leu	Ser
	130					135					140				
Ala	Ser	Val	Gly	Asp	Arg	Val	Thr	Ile	Thr	Cys	Arg	Ala	Ser	Gln	Gly
145					150					155					160
Ile	Ser	Ser	Tyr	Leu	Ala	Trp	Tyr	Gln	Gln	Lys	Pro	Gly	Lys	Ala	Pro
				165					170					175	
Lys	Leu	Leu	Val	Tyr	Ala	Ala	Ser	Thr	Leu	Gln	Ser	Gly	Val	Pro	Ser
			180					185					190		
Arg	Phe	Ser	Gly	Ser	Gly	Ser	Gly	Thr	Glu	Phe	Thr	Leu	Thr	Ile	Ser
		195					200					205			
Ser	Leu	Gln	Pro	Glu	Asp	Phe	Ala	Thr	Tyr	Tyr	Cys	Gln	Gln	Leu	Asn
	210					215					220				
Ser	Tyr	Arg	Leu	Thr	Phe	Gly	Gln	Gly	Thr	Lys	Leu	Glu	Ile	Lys	Arg
225					230					235					240

<210> 14
 <211> 742
 <212> DNA
 <213> Homo sapiens

<400> 14
 agcttcagga tcttgaaagg ttttgctcta cttcctgaag acctgaacac cgctcccata 60
 aagccatggc ttgccttgga tttcagcggc acaaggctca gctgaacctg gctaccagga 120
 cctggccctg cactctcctg ttttttcttc tcttcatccc tgtcttctgc aaagcaatgc 180
 acgtggccca gcctgctgtg gtactggcca gcagccgagg catcgccagc tttgtgtgtg 240
 agtatgcac tccaggcaaa gccactgagg tccgggtgac agtgcttcgg caggctgaca 300
 gccaggtgac tgaagtctgt gcggcaacct acatgatggg gaatgagttg accttcctag 360
 atgattccat ctgcacgggc acctccagtg gaaatcaagt gaacctcact atccaaggac 420
 tgagggccat ggacacggga ctctacatct gcaaggtgga gctcatgtac ccaccgccat 480
 actacctggg cataggcaac ggaacccaga tttatgtaat tgatccagaa ccgtgcccag 540
 attctgactt cctcctctgg atccttgacag cagttagttc ggggttggtt ttttatagct 600
 ttctcctcac agctgtttct ttgagcaaaa tgctaaagaa aagaagccct cttacaacag 660
 gggcttatgt gaaaatgccc ccaacagagc cagaatgtga aaagcaattt cagccttatt 720
 ttattcccat caattgagaa tt 742

<210> 15
 <211> 223
 <212> PRT
 <213> Homo sapiens

<400> 15
 Met Ala Cys Leu Gly Phe Gln Arg His Lys Ala Gln Leu Asn Leu Ala
 1 5 10 15
 Thr Arg Thr Trp Pro Cys Thr Leu Leu Phe Phe Leu Leu Phe Ile Pro
 20 25 30
 Val Phe Cys Lys Ala Met His Val Ala Gln Pro Ala Val Val Leu Ala
 35 40 45
 Ser Ser Arg Gly Ile Ala Ser Phe Val Cys Glu Tyr Ala Ser Pro Gly
 50 55 60
 Lys Ala Thr Glu Val Arg Val Thr Val Leu Arg Gln Ala Asp Ser Gln
 65 70 75 80
 Val Thr Glu Val Cys Ala Ala Thr Tyr Met Met Gly Asn Glu Leu Thr
 85 90 95
 Phe Leu Asp Asp Ser Ile Cys Thr Gly Thr Ser Ser Gly Asn Gln Val
 100 105 110
 Asn Leu Thr Ile Gln Gly Leu Arg Ala Met Asp Thr Gly Leu Tyr Ile
 115 120 125

Cys Lys Val Glu Leu Met Tyr Pro Pro Pro Tyr Tyr Leu Gly Ile Gly
 130 135 140
 Asn Gly Thr Gln Ile Tyr Val Ile Asp Pro Glu Pro Cys Pro Asp Ser
 145 150 155 160
 Asp Phe Leu Leu Trp Ile Leu Ala Ala Val Ser Ser Gly Leu Phe Phe
 165 170 175
 Tyr Ser Phe Leu Leu Thr Ala Val Ser Leu Ser Lys Met Leu Lys Lys
 180 185 190
 Arg Ser Pro Leu Thr Thr Gly Val Tyr Val Lys Met Pro Pro Thr Glu
 195 200 205
 Pro Glu Cys Glu Lys Gln Phe Gln Pro Tyr Phe Ile Pro Ile Asn
 210 215 220

<210> 16
 <211> 773
 <212> DNA
 <213> Homo sapiens

<400> 16
 aagcttcgag ccaagcagcg tectggggag cgcgtcatgg ccttaccagt gaccgccttg 60
 ctectgccgc tggccttgct gctccacgcc gccaggccga gccagttccg ggtgtcgccg 120
 ctggatcgga cctggaacct gggcgagaca gtggagctga agtgccaggt gctgctgtcc 180
 aaccgcagct cgggctgctc gtggctcttc cagccgcgcg gcgccgccgc cagtcccacc 240
 ttctctctat acctctccca aaacaagccc aaggcggccg aggggctgga caccagcgcg 300
 ttctcgggca agaggttggg ggacaccttc gtcctcaccg tgagcgactt ccgccgagag 360
 aacgagggct actatttctg ctgcggcctg agcaactcca tcatgtactt cagccacttc 420
 gtgccggtct tectgccagc gaagcccacc acgacgccag cgccgcgacc accaacaccg 480
 gcgccaccca tcgcgtcgca gccctgtgct ctgcgcccag aggcgtgccg gccagcgggc 540
 gggggcgagc tgcacacgag ggggctggac ttgcctgtg atatctacat ctggggcgccc 600
 ttggccggga cttgtggggc ctttctctctg tcaactggta tcacccttta ctgcaaccac 660
 aggaaccgaa gacgtgtttg caaatgtccc cggcctgtgg tcaaatcggg agacaagccc 720
 agcctttcgg cgagatacgt ctaaccctgt gcaacagcca ctacatgaat tcc 773

<210> 17
 <211> 28
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> PCR primer

<400> 17
 ttgaagctta gccatggctt gctctgga

<210> 18
 <211> 33
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> PCR primer

<400> 18
 taatgaattc tcaattgatg ggaataaaat aag 33

<210> 19
 <211> 60
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> PCR primer

<400> 19
 cgggttctgca gcaccaccgg agccaccatc agaatctggg catggttctg gatcaatgac 60

<210> 20
 <211> 25
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> PCR primer

<400> 20
 gagctgaaac gggcggccgc agaac 25

<210> 21
 <211> 22
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> PCR primer

<400> 21
 ctggcctgca gcattcagat cc 22

<210> 22
 <211> 30
 <212> DNA
 <213> Artificial Sequence

<220>

<223> PCR primer

<400> 22

ttcaaagctt caggatcctg aaagggttttg

30

<210> 23

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> PCR primer

<400> 23

taatgaattc tcaattgatg ggaataaaaat aag

33

<210> 24

<211> 76

<212> DNA

<213> Artificial Sequence

<220>

<223> PCR primer

<400> 24

gatgtagata tcacaggcga agtcgacacc accggagcca ccaattacat aaatctgggc 60
tccgttgect atgccc 76

<210> 25

<211> 29

<212> DNA

<213> Artificial Sequence

<220>

<223> PCR primer

<400> 25

tcgcgcccga gcttcgagcc aagcagcgt

29

<210> 26

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> PCR primer

<400> 26

taatgaattc tcaattgatg ggaataaaaat aag

33

<210> 27

<211> 73

<212> DNA

<213> Artificial Sequence

<220>

<223> PCR primer

<400> 27

cataggcaac ggagcccaga tttatgtaat tgggtggctcc ggtgggtgtcg acttcgcctg 60
tgatatctac atc 73